

Professor Joshua Lederberg

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Department of Genetics

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Dear Professor Lederberg:

I have received your reprints on 'transduction' for which I am exceedingly grateful. I am greatly interested in your papers, since I and My associates have been studying induction of antigenic changes in the structure of O antigens of Salmonella E<sub>1</sub>, E<sub>2</sub> and E<sub>z</sub> group, and have observed recently a phenomenon similar to transduction. As the details of the data have not been published as yet, important points are described in the enclosed papers.

I was thinking that you might be interested in the results of our experiments. I would appreciate your opinions and suggestions if I am not asking too much . .

Yours sincerely,

*Hisao Uetake*

Hisao Uetake, M.D.

Professor

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It has been found that the organisms of Salmonella E<sub>2</sub> group are lysogenic and the bacteriophage obtained from them are capable of infecting the organisms of E<sub>1</sub> group. When the organisms of E<sub>1</sub> group were exposed to the phages, some of them were lysogenized and simultaneously, in lysogenized cells, their O antigens underwent a change from III,X to III,XV.

S.newington, S.selandia, S.new-brunswick, S.cambridge, S.kinshasa, S.canoga, and S.thomasville of E<sub>2</sub> group have been found to be lysogenic.

S.london, S.give, S.anatum, S.amager, S.zanzibar, S.shangani, S.batantan, S.vejle, S.meleagridis, S.elizabethville, S.simi, S.weltevreden and S.orion of E<sub>1</sub> group have all been found to be receptive to the phage and to be lysogenized, resulting in the antigenic changes from III,X to III,XV. O antigens of induced variants reverted to the original structure by cultivating the variant strains in broth containing XV antiserum, which was prepared by absorbing S.newington O antiserum with S.anatum.

In S.lexington and S.macallen, O antigens were changed from III,X to III,X,XV, which have not been observed as yet in naturally isolated Salmonella strains.

In S.chittagon O antigens were changed from I,III,X,XIX to III,XV, which reverted to I,III,X,XIX by cultivating the variant in broth containing XV antiserum, prepared as described above.

In S.niloese and S.senftenberg-simsbury, O antigens were altered by the phage from I,III,XIX to I,III,XIX,XV, which reverted to the original structure by cultivating the variants

in broth containing XV antiserum, prepared as described above.

In S.senftenberg 87Aa', which was a variant of S.senftenberg A and has only III as its major O antigen, O antigenic structure was converted from III to III,XV by the phage of S.canoga.

The above-mentioned findings indicate that strains of E<sub>1</sub> group or E<sub>3</sub> group can be lysogenized by infection with phage, which are obtained from lysogenic strains of E<sub>2</sub> group, and simultaneously their O antigenic structures are changed from III,X to III, XV or from I,III, XIX to I,III,XIX,XV and or in the special case of S. senftenberg 87Aa' from III to III,XV. It is noteworthy that the changes of O antigenic structure are always associated with lysogenization of the cultures. The induced variants, which contain XV antigenic factor as a part of O antigens, are invariably lysogenic and the phages obtained from them are capable of inducing the same antigenic variation in organisms of E<sub>1</sub> group as those from strains of E<sub>2</sub> group.

The activities inducing antigenic changes were not inactivated by desoxyribonuclease, but they were inactivated when the phages were inactivated.

These findings seem to be similar to Groman's findings on the relationship between toxigenicity and lysogenicity of *Corynebacterium diphtheriae*.

The followings are differences between your observations and ours.

- 1) You have observed that phage is not always transmitted to all the progeny of cells showing transduced character, but in

our experiments formation of XV antigenic factor is always associated with establishment of lysogenicity as far as tested.

2) In your experiments transducing lysates were prepared by adding phage 22 to a freshly seeded broth culture of the donor strain. In our experiments phage preparations were obtained by autolysis of cultures of E<sub>2</sub> group or by filtration of mixed broth cultures of E<sub>1</sub> and E<sub>2</sub> group cells. Even when phages were propagated on cells of E<sub>1</sub> group, they possessed activities inducing formation of XV antigenic factor. In so far as the formation of XV antigenic factor was concerned, the activity of the phage seemed not to be affected by strains on which phage was propagated.

3) In transduction you have pointed out replacements of genetic traits, while in our experiments there has been no evidence of such a replacement as yet.